(FILE 'HOME' ENTERED AT 12:16:23 ON 03 NOV 2005)

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FILE'AGRICOLA, MEDLINE, CAPLUS, BIOSIS' ENTERED AT 12:16:26 ON 03 NOV
2005L111349 S (NEUTRAL (1N) ENDOPEPTIDASE) OR NEP OR NEPRILYSINL2900 S L1 AND (MUS OR MOUSE)L3323 S L2 AND (CDNA OR CLON? OR GENE)L4105 S L3 AND PY<1999</td>L569 DUP REM L4 (36 DUPLICATES REMOVED)L60 S L5 AND 765

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- L6 ANSWER 10 OF 12 CAPLUS COPYRIGHT 2005 ACS on STN
- AN 1993:186782 CAPLUS
- DN 118:186782

TI Murine common acute lymphoblastic leukemia antigen (CD10 neutral endopeptidase 24.11). Molecular characterization, chromosomal localization, and modeling of the active site

- AU Chen, Chang Yan; Salles, Gilles; Seldin, Michael F.; Kister, Alexander E.; Reinherz, Ellis L.; Shipp, Margaret A.
- CS Lab. Immunobiol., Harvard Med. Sch., Boston, MA, 02115, USA
- SO Journal of Immunology (**1992**), 148(9), 2817-25
- CODEN: JOIMA3; ISSN: 0022-1767
- DT Journal
- LA English
- AB To further analyze antigen CD10/neutral endopeptidase 24.11 CD10/NEP] function in lymphoid and nonlymphoid cells using well characterized murine systems, the murine CD10/NEP homolog was isolated, its chromosomal location was determined and the enzyme active site was modeled. The murine CD10/NEP cDNA predicts a 750-amino acid (aa) type II integral membrane protein with 90% identity to the human CD10 sequence and 100% conservation of critical aa and functional motifs. The latter include the pentapeptide consensus sequence required for zinc binding and catalytic activity, addnl. aa associated with substrate binding, and the extracellular cysteines that participate in disulfide bonds required for enzymic activity. Like its human homolog, murine CD10/NEP has multiple alternative 5'-untranslated region sequences. The gene is localized on the proximal half of murine chromosome 3. In Northern anal., murine CD10/NEP transcripts are abundant in bone marrow stromal cells that support pre-B cell differentiation but are undetectable in representative Abelson transformed pre-B cell lines. The murine CD10/NEP active site was modeled by aligning critical conserved CD10/NEP residues with comparable residues in the active site of thermolysin, a bacterial metalloprotease with similar substrate specificity. The model predicts that the 2 enzymes have similar clefts that comprise the active site and permit zinc-dependent substrate interactions.

ANSWER 8 OF 12 L6 MEDLINE on STN

DUPLICATE 4

- AN 93390947 MEDLINE DN
- PubMed ID: 8397369
- ΤI NEP: a novel receptor-like tyrosine kinase expressed in proliferating neuroepithelia.
- AU Zerlin M; Julius M A; Goldfarb M
- CS Department of Biochemistry and Molecular Biophysics, Columbia University College of Physicians and Surgeons, New York, New York 10032.
- SO Oncogene, (1993 Oct) 8 (10) 2731-9.
- Journal code: 8711562. ISSN: 0950-9232.
- CY ENGLAND: United Kingdom
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- Priority Journals FS
- ΕM 199310
- Entered STN: 19931105 ED Last Updated on STN: 19931105 Entered Medline: 19931018

We have isolated a murine **cDNA**, nep, which encodes a novel AB receptor-like protein tyrosine kinase. The kinase region of NEP protein bears 50% amino acid sequence identity to the neurotrophin receptors (TRKs). While the intracytoplasmic portion of NEP also contains a short kinase insert region and C-terminal tail reminiscent of the TRK proteins, the putative extracellular domain of NEP is unrelated to any known proteins. The nep gene is strongly expressed within proliferating neuroepithelia of mouse embryos, commencing at the early somite stage (embryonic day 8.0) and persisting in the proliferative ventricular zones of the brain and spinal cord, suggesting that one function of NEP kinase is to signal proliferation of neuroepithelial cells in response to an as yet unknown ligand. The nep gene is also expressed in embryonic sensory ganglia, striated muscle and epidermis, as well as in several adult tissues, including the ventricle linings and glia subpopulations in the brain.

Neutral endopeptidase modulation of septic shock. AU Lu B; Gerard N P; Kolakowski L F Jr; Bozza M; Zurakowski D; Finco O; Carroll M C; Gerard C CS Ina Sue Perlmutter Laboratory, Children's Hospital, Boston, Massachusetts, USA. NC HL19170 (NHLBI) HL51366 (NHLBI) Journal of experimental medicine, (1995 Jun 1) 181 (6) 2271-5. SO Journal code: 2985109R. ISSN: 0022-1007. CY United States DTJournal; Article; (JOURNAL ARTICLE) LA English FS Priority Journals EΜ 199506 Entered STN: 19950707 ED Last Updated on STN: 19950707 Entered Medline: 19950623 AB Neutral endopeptidase (NEP; EC. 3.4.24.11) is a type 2 cell surface metalloprotease known by a variety of eponyms, including enkephalinase, common acute lymphoblastic leukemia antigen, and CD10. Identified substrates are largely neural or humoral oligopeptide agonists, and the enzyme functions to terminate signaling by degrading the ligand, analogously to acetylcholine/acetylcholinesterase. Targeted disruption of the NEP locus in mice results in enhanced lethality to endotoxin shock with a pronounced gene dosage effect. The site(s) of action appears downstream from release of tumor necrosis factor and interleukin-1 since NEP-deficient animals demonstrate increased

sensitivity to these mediators as well. This unexpected finding indicates

an important protective role for NEP in septic shock.

1 × 1

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Trp Thr Leu Phe Ile Gln Asn Val Leu Ser Ser Val Glu Val Glu Leu Phe Pro Asp Glu Glu Val Val Val Tyr Gly Ile Pro Tyr Leu Glu Asn Leu Glu Asp Ile Ile Asp Ser Tyr Ser Ala Arg Thr Met Gln Asn Tyr Leu Val Trp Arg Leu Val Leu Asp Arg Ile Gly Ser Leu Ser Gln Arg Phe Lys Glu Ala Arg Val Asp Tyr Arg Lys Ala Leu Tyr Gly Thr Thr Val Glu Glu Val Arg Trp Arg Glu Cys Val Ser Tyr Val Asn Ser Asn Met Glu Ser Ala Val Gly Ser Leu Tyr Ile Lys Arg Ala Phe Ser Lys Asp Ser Lys Ser Thr Val Arg Glu Leu Ile Glu Lys Ile Arg Ser Val Phe Val Asp Asn Leu Asp Glu Leu Asn Trp Met Asp Glu Glu Ser Lys Lys Lys Ala Gln Glu Lys Ala Met Asn Ile Arg Glu Gln Ile Gly Tyr Pro Asp Tyr Ile Leu Glu Asp Asn Asn Lys His Leu Asp Glu Glu Tyr Ser Ser Leu Thr Phe Tyr Glu Asp Leu Tyr Phe Glu Asn Gly Leu Gln Asn Leu Lys Asn Asn Ala Gln Arg Ser Leu Lys Lys Leu Arg Glu Lys Val Asp Gln Asn Leu Trp Ile Ile Gly Ala Ala Val Val Asn Ala Phe Tyr Ser Pro Asn Arg Asn Gln Ile Val Phe Pro Ala Gly Ile Leu Gln Pro Pro Phe Phe Ser Lys Asp Gln Pro Gln Ser Leu Asn Phe Gly Gly Ile Gly Met Val Ile Gly His Glu Ile Thr His Gly Phe Asp Asp Asn Gly Arg Asn Phe Asp Lys Asn Gly Asn Met Leu Asp Trp Trp Ser Asn Phe Ser Ala Arg His Phe Gln Gln Gln Ser Gln Cys Met Ile Tyr Gln

Tyr Gly Asn Phe Ser Trp Glu Leu Ala Asp Asn Gln Asn Val Asn Gly Phe Ser Thr Leu Gly Glu Asn Ile Ala Asp Asn Gly Gly Val Arg Gln Ala Tyr Lys Ala Tyr Leu Arg Trp Leu Ala Asp Gly Gly Lys Asp Gln Arg Leu Pro Gly Leu Asn Leu Thr Tyr Ala Gln Leu Phe Phe Ile Asn Tyr Ala Gln Val Trp Cys Gly Ser Tyr Arg Pro Glu Phe Ala Val Gln Ser Ile Lys Thr Asp Val His Ser Pro Leu Lys Tyr Arg Val Leu Gly Ser Leu Gln Asn Leu Pro Gly Phe Ser Glu Ala Phe His Cys Pro Arg Gly Ser Pro Met His Pro Met Lys Arg Cys Arg Ile Trp <210> 14 <211> 2676 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (7)..(2316) <400> 14 gtgggg atg gtg gag agc gcc ggc cgt gca ggg cag aag cgc ccg ggg Met Val Glu Ser Ala Gly Arg Ala Gly Gln Lys Arg Pro Gly ttc ctg gag ggg ggg ctg ctg ctg ctg ctg ctg gtg acc gct gcc Phe Leu Glu Gly Gly Leu Leu Leu Leu Leu Leu Leu Val Thr Ala Ala ctg gtg gcc ttg ggt gtc ctc tac gcc gac cgc aga ggg aag cag ctg Leu Val Ala Leu Gly Val Leu Tyr Ala Asp Arg Arg Gly Lys Gln Leu cca cgc ctt gct agc cgg ctg tgc ttc tta cag gag gag agg acc ttt Pro Arg Leu Ala Ser Arg Leu Cys Phe Leu Gln Glu Glu Arg Thr Phe gta aaa cga aaa ccc cga ggg atc cca gag gcc caa gag gtg agc gag Val Lys Arg Lys Pro Arg Gly Ile Pro Glu Ala Gln Glu Val Ser Glu